STAFF USE ONLY	
	SYSTEMS 1
	DARC/QUESTEL
TIME	DIALOG
	SDC
	STAFF USE ONLY

=> file caplus
FILE 'CAPLUS' ENTERED AT 15:29:05 ON 18 AUG 2003
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FILE COVERS 1907 - 18 Aug 2003 VOL 139 ISS 8 FILE LAST UPDATED: 17 Aug 2003 (20030817/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> d que 117

```
1 SEA FILE=REGISTRY ABB=ON PLU=ON EPI-2-INOSOSE/CN
               24 SEA FILE=CAPLUS ABB=ON PLU=ON L13
L16
                1 SEA FILE=CAPLUS ABB=ON PLU=ON L16(L)(BPN OR BMF)/RL & any preparation of epi-2-1
in usose by biological meuro
(micro organism)
L17
=> d que 126
                1 SEA FILE=REGISTRY ABB=ON PLU=ON EPI-2-INOSOSE/CN
L15
                1 SEA FILE=REGISTRY ABB=ON PLU=ON EPI-INOSITOL/CN
L16
               24 SEA FILE=CAPLUS ABB=ON PLU=ON L13
L23
              102 SEA FILE=CAPLUS ABB=ON PLU=ON L15
               10 SEA FILE=CAPLUS ABB=ON PLU=ON LIS

10 SEA FILE=CAPLUS ABB=ON PLU=ON L23(L)PREP/RL & prep (any memo) of epi-inosity)

7 SEA FILE=CAPLUS ABB=ON PLU=ON L16(L)(RCT OR RACT)/RL & epi-2-inosose as a react-
L24
L25
                1 SEA FILE=CAPLUS ABB=ON PLU=ON L24 AND L25 ( ci+e
L26
```

=> s 117 or 126

1 L17 OR L26 1 cite (applicant)

=> file casreact

FILE 'CASREACT' ENTERED AT 15:29:08 ON 18 AUG 2003 USE IS SUBJECT TO THE TERMS OF YOUR CUSTOMER AGREEMENT COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY (ACS)

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FILE CONTENT:1907 - 17 Aug 2003 VOL 139 ISS 7

Some records from 1974 to 1991 are derived from the ZIC/VINITI data file and provided by InfoChem and some records are produced using some INPI data from the period prior to 1986.

This file contains CAS Registry Numbers for easy and accurate substance identification.

Crossover limits have been increased. See HELP RNCROSSOVER for details.

Structure search limits have been raised. See HELP SLIMIT for the new, higher limits.

```
=> d que 136
```

```
1 SEA FILE=CASREACT ABB=ON PLU=ON 6623-68-3/PRO - prep of epi-2-in 0 sole any 2 SEA FILE=CASREACT ABB=ON PLU=ON (L34 OR L35) 2 Cites epi-1n 0 site means
```

=> file uspatful

FILE 'USPATFULL' ENTERED AT 15:29:09 ON 18 AUG 2003
CA INDEXING COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY (ACS)

FILE COVERS 1971 TO PATENT PUBLICATION DATE: 14 Aug 2003 (20030814/PD)
FILE LAST UPDATED: 14 Aug 2003 (20030814/ED)
HIGHEST GRANTED PATENT NUMBER: US6606748
HIGHEST APPLICATION PUBLICATION NUMBER: US2003154532
CA INDEXING IS CURRENT THROUGH 14 Aug 2003 (20030814/UPCA)
ISSUE CLASS FIELDS (/INCL) CURRENT THROUGH: 14 Aug 2003 (20030814/PD)
REVISED CLASS FIELDS (/NCL) LAST RELOADED: Jun 2003
USPTO MANUAL OF CLASSIFICATIONS THESAURUS ISSUE DATE: Jun 2003

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> d que 170

L 1 4	1 SEA FILE=REGISTRY ABB=ON PLU=ON MYO-INOSITOL/CN
LS8	31 SEA FILE=USPATFULL ABB=ON PLU=ON ?INOSOSE
L59	1910 SEA FILE=USPATFULL ABB=ON PLU=ON MYO-INOSITOL OR L14
L65	7 SEA FILE=USPATFULL ABB=ON PLU=ON L58(P)L59
L66	1 SEA FILE=USPATFULL ABB=ON PLU=ON L65(P)(MICROB? OR MICROORG?)
L67	4 SEA FILE=USPATFULL ABB=ON PLU=ON L65(P)(OXIDI? OR OXIDA?)
L68	3 SEA FILE=USPATFULL ABB=ON PLU=ON L65 AND (PSEUDOMONAS OR
	XANTHOMONAS OR ACETOBACTER OR GLUCONOBACTER OR AGROBACTER? OR
	ERWINIA OR ENTEROBACTER OR SERRATIA OR YERSINIA OR PASTEURELLA
	OR HAEMOPHIL?)
L69	O SEA FILE=USPATFULL ABB=ON PLU=ON L65 AND FERM(W)BP(W)(7168
	OR 7170 OR 7169 OR 10135 OR 10215 OR 10119)
L70	5 SEA FILE=USPATFULL ABB=ON PLU=ON (L66 OR L67 OR L68 OR L69) 5 cites
	2 (and an en

=> d que 177

L15	1 SEA FILE=REGISTRY ABB=ON PLU=ON EPI-INOSITOL/CN
L58	31 SEA FILE=USPATFULL ABB=ON PLU=ON ?INOSOSE
L60	49 SEA FILE=USPATFULL ABB=ON PLU=ON EPI-INOSITOL OR L15
L76	3 SEA FILE=USPATFULL ABB=ON PLU=ON (REDUC? OR BOROHYDRID?) AND
	L58 AND L60
L77	2 SEA FILE=USPATFULL ABB=ON PLU=ON L76 NOT VANADIUM/TI 2 CITES

=> s 170 or 177

L82 6 L70 OR L77 6 patents

=> file scisearch

FILE 'SCISEARCH' ENTERED AT 15:29:12 ON 18 AUG 2003 COPYRIGHT 2003 THOMSON ISI

FILE COVERS 1974 TO 15 Aug 2003 (20030815/ED)

=> d que 156 149 30 SEA FILE=SCISEARCH ABB=ON PLU=ON EPI-INOSITOL L55 59 SEA FILE=SCISEARCH ABB=ON PLU=0N ?INOSOSE 2 SEA FILE=SCISEARCH ABB=ON PLU=ON L55 AND L49 2 cites Sci Search L56 => file wpix FILE 'WPIX' ENTERED AT 15:29:13 ON 18 AUG 2003 COPYRIGHT (C) 2003 THOMSON DERWENT FILE LAST UPDATED: 13 AUG 2003 <20030813/UP> MOST RECENT DERWENT UPDATE: <200352/DW> 200352 DERWENT WORLD PATENTS INDEX SUBSCRIBER FILE, COVERS 1963 TO DATE >>> NEW WEEKLY SDI FREQUENCY AVAILABLE --> see NEWS <<< >>> SLART (Simultaneous Left and Right Truncation) is now available in the /ABEX field. An additional search field /BIX is also provided which comprises both /BI and /ABEX <<< >>> PATENT IMAGES AVAILABLE FOR PRINT AND DISPLAY <<< >>> FOR DETAILS OF THE PATENTS COVERED IN CURRENT UPDATES, SEE http://www.derwent.com/dwpi/updates/dwpicov/index.html <<< >>> FOR A COPY OF THE DERWENT WORLD PATENTS INDEX STN USER GUIDE, PLEASE VISIT: http://www.stn-international.de/training_center/patents/stn_guide.pdf <<< >>> FOR INFORMATION ON ALL DERWENT WORLD PATENTS INDEX USER GUIDES, PLEASE VISIT: http://www.derwent.com/userguides/dwpi_guide.html <<< => d que 139 245 SEA FILE=WPIX ABB=ON PLU=ON MYO-INOSITOL L37 2 SEA FILE=WPIX ABB=ON PLU=ON EPI-2-INOSOSE L38 1 SEA FILE=WPIX ABB=ON PLU=ON L37 AND L38 139 => d que 141 2 SEA FILE=WPIX ABB=ON PLU=ON EPI-2-INOSOSE L38 10 SEA FILE=WPIX ABB=ON PLU=ON EPI-INOSITOL L40 2 SEA FILE=WPIX ABB=ON PLU=ON L40 AND L38 L41 => d que 142 2 SEA FILE=WPIX ABB=ON PLU=ON EPI-2-INOSOSE L38 10 SEA FILE=WPIX ABB=ON PLU=ON EPI-INOSITOL L40 L41 2 SEA FILE=WPIX ABB=ON PLU=ON L40 AND L38 1 SEA FILE=WPIX ABB=ON PLU=ON L41 AND BOROHYDRIDE L42 => s 139 or 141-42 2 cites WPIX 2 L39 OR (L41 OR L42)

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Searched by Susan Hanley 305-4053

duplicates

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FILE 'CASREACT' ENTERED AT 15:30:17 ON 18 AUG 2003
USE IS SUBJECT TO THE TERMS OF YOUR CUSTOMER AGREEMENT
COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY (ACS)
FILE 'USPATFULL' ENTERED AT 15:30:17 ON 18 AUG 2003
CA INDEXING COPYRIGHT (C) 2003 AMERICAN CHEMICAL SOCIETY (ACS)
FILE 'SCISEARCH' ENTERED AT 15:30:17 ON 18 AUG 2003
COPYRIGHT 2003 THOMSON ISI
FILE 'WPIX' ENTERED AT 15:30:17 ON 18 AUG 2003
COPYRIGHT (C) 2003 THOMSON DERWENT
PROCESSING COMPLETED FOR L81
PROCESSING COMPLETED FOR L36
PROCESSING COMPLETED FOR L82
PROCESSING COMPLETED FOR L56
PROCESSING COMPLETED FOR L83
             10 DUP REM L81 L36 L82 L56 L83 (3 DUPLICATES REMOVED) /O
                                                                          cites total
184
                ANSWER '1' FROM FILE CAPLUS
                ANSWER '2' FROM FILE CASREACT
                ANSWERS '3-8' FROM FILE USPATFULL
                ANSWER '9' FROM FILE SCISEARCH
                ANSWER '10' FROM FILE WPIX
=> d ibib abs hitstr 1
L84 ANSWER 1 OF 10 CAPLUS COPYRIGHT 2003 ACS on STN DUPLICATE 1
                         2000:881342 CAPLUS
ACCESSION NUMBER:
DOCUMENT NUMBER:
                         134:42384
TITLE:
                         Novel process for producing L-epi-2-inosose by
                         microbial oxidation of myo-inositol and novel process
                         for producing epi-inositol
INVENTOR(S):
                         Takahashi, Atsushi; Kanbe, Kenji; Mori, Tetsuya; Kita,
                         Yuichi; Tamamura, Tsuyoshi; Takeuchi, Tomio
                         Hokko Chemical Industry Co., Ltd., Japan; Zaidan Hojin
PATENT ASSIGNEE(S):
                         Biseibutsu Kagaku Kenkyu Kai
SOURCE:
                         PCT Int. Appl., 65 pp.
                         CODEN: PIXXD2
DOCUMENT TYPE:
                         Patent
                         Japanese
LANGUAGE:
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
     PATENT NO.
                      KIND DATE
                                           APPLICATION NO.
                                                            DATE
                            20001214
                                           WO 2000-JP3687
                                                            20000607
     WO 2000075355
                      A1
         W: CA, CN, IL, IN, JP, KR, US
         RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
             PT, SE
                                           EP 2000-937174
     EP 1197562
                            20020417
                                                            20000607
                       A1
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, SI, LT, LV, FI, RO
PRIORITY APPLN. INFO.:
                                        JP 1999-159861
                                                            19990607
                                                         Α
                                        JP 1999-340523
                                                            19991130
                                        JP 2000-151709
                                                            20000523
                                        WO 2000-JP3687
                                                            20000607
OTHER SOURCE(S):
                         CASREACT 134:42384
    L-Epi-2-inosose and epi-inositol, which are useful as various drugs or
     synthesis intermediates, can be efficiently produced from less expensive
     myo-inositol. Myo-inositol is treated with a gram-neg. bacterium. e.g.
    Xanthomonas sp., capable of converting myo-inositol into L-epi-2-inosose
```

to thereby convert the myo-inositol into L-epi-2-inosose. The

L-epi-2-inosose thus obtained is further reacted in an aq. reaction medium with a reducing agent comprising an alkali metal boron hydride or another alkali metal hydride to form epi-inositol and myo-inositol. Next, the epi-inositol is sepd. and isolated from the redn. reaction mixt.

comprising epi-inositol and myo-inositol to give epi-inositol.

6623-68-3P, epi-2-Inosose

RL: BPN (Biosynthetic preparation); RCT (Reactant);

BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent)

(novel process for producing L-epiinosose by microbial oxidn. of

myo-inositol and boron hydride-redn. to epi-inositol)

6623-68-3 CAPLUS RN

CN epi-2-Inosose (9CI) (CA INDEX NAME)

Relative stereochemistry.

488-58-4P, epi-Inositol

RL: SPN (Synthetic preparation); PREP (Preparation)

(novel process for producing L-epiinosose by microbial oxidn. of

myo-inositol and boron hydride-redn. to epi-inositol)

488-58-4 CAPLUS RN

epi-Inositol (9CI) (CA INDEX NAME)

Relative stereochemistry.

REFERENCE COUNT:

THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

=> d ibib abs fcrdref 2

L84 ANSWER 2 OF 10 CASREACT COPYRIGHT 2003 ACS on STN DUPLICATE 2

7

ACCESSION NUMBER:

133:105232 CASREACT

TITLE:

Rare and complex saccharides from D-galactose and other milk-derived carbohydrates. Part 12. A new highly diastereoselective synthesis of epi-inositol

from D-galactose

AUTHOR(S):

Pistara, Venerando; Barili, Pier Luigi; Catelani, Giorgio; Corsaro, Antonino; D'Andrea, Felicia;

Fisichella, Salvatore

CORPORATE SOURCE:

Dipartimento di Scienze Chimiche, Universita degli

Studi di Catania, Catania, I-95125, Italy Tetrahedron Letters (2000), 41(17), 3253-3256

CODEN: TELEAY; ISSN: 0040-4039

PUBLISHER:

Elsevier Science Ltd.

DOCUMENT TYPE:

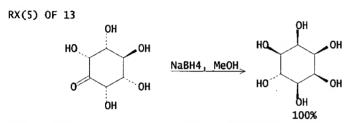
Journal

LANGUAGE:

SOURCE:

English

The inosose deriv. I (Bn = PhCH2) was obtained with high stereoselectivity by intramol. aldol condensation of the aldohexos-5-ulose II, and it was selectively reduced and debenzylated to give epi-inositol in high yield. The stereochem, and the preferred conformations of the compds. were detd. through 1D- and 2D-NMR expts.



REF: Tetrahedron Letters, 41(17), 3253-3256; 2000 NOTE: stereoselective

REFERENCE COUNT:

THERE ARE 23 CITED REFERENCES AVAILABLE FOR THIS 23 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

=> d ibib abs hitrn kwic 3-8

L84 ANSWER 3 OF 10 USPATFULL on STN

ACCESSION NUMBER: 2002:88660 USPATFULL

TITLE:

Labelled phosphoinositides and analogues Aneja, Rajindra, Ithaca, NY, United States INVENTOR(S):

PATENT ASSIGNEE(S): Nutrimed Biotech, Ithaca, NY, United States (U.S.

corporation)

•				
	NUMBER	KIND	DATE	
PATENT INFORMATION:	US 6376697	B1		
APPLICATION INFO.:	US 1999-292242		19990415	(9)
	NUMBER	DA	TE	
PRIORITY INFORMATION: DOCUMENT TYPE:	US 1998-81847P Utility	1998	0415 (60)	
FILE SEGMENT: PRIMARY EXAMINER:	GRANTED Ambrose, Michael	G.		
LEGAL REPRESENTATIVE:	Williams, Morgan and Amerson			
NUMBER OF CLAIMS: EXEMPLARY CLAIM:	29 1			
NUMBER OF DRAWINGS: LINE COUNT:	0 Drawing Figure 1102	(s); 0	Drawing Pa	ige(s)
LINE COURT	1105			

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention provides novel compounds comprising cellular phosphoinositides and analogues tagged with stable or radioactive isotopes. The present invention also provides novel methods for the preparation of the said phosphoinositides by syntheses, and novel key intermediates of synthesis; the novel methods of synthesis are applied also for the preparation of the phosphoinositides in non-labelled form. In addition, the present invention discloses a class of novel compounds as isotope labelled key precursors of labelled phosphoinositides. These precursors are derivatives of the target phosphoinositides, labelled with stable or radioactive isotopes, wherein OH and phosphate groups are blocked with temporary protecting groups.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

- DETD In the second approach, outlined in Scheme 2, a selectively protected myo-inositol, e.g., 1D-2,6-di-O-benzyl-myo-inositol-3,4,5-tris(dibenzylphosphate) 10, wherein only the equatorial 1-OH is unprotected, and all other OH groups are blocked with temporary protecting groups is a suitable starting material. Oxidation of 10 (Step a) to the corresponding inosose 11 is carried out using the reagent mixture comprising dimethylsulfoxide and acetic anhydride (DMSO-Ac.sub.20). A hydrogen, deuterium or tritium atom is introduced (Step b) by reduction of inosose 11, using NaBH.sub.4, NaB.sup.2H.sub.4 or NaB.sup.3H.sub.4, to the corresponding secondary alcohol 12 carrying H--C--OH, .sup.2H--C--OH or .sup.3H--C--OH labels. The product. . . pyridine. The purified product 13 is the labelled precursor analogous with 3, and is deprotected by H.sub.2-Pd/C to labelled 1D-1-(1',2'-O-dipalmitoyl -sn-glycero-3'-phospho)-myo-inositol-3,4,5-trisphosphate (DPPtdIns-3,4,5-P.sub.3) 14. ##STR5##
- The phosphatidyl-inosose (7) and inosose (11) DETD employed in Scheme 1 and 2 respectively are important novel intermediates. Equally useful are the 2-phosphatidyl-1-keto and 1-phosphatidyl-6-keto structural isomers of 7 and the 2-keto and 6-keto isomers of 11 prepared by oxidation of the corresponding 1-phosphatidyl-1-OH and 1-phosphatidyl-6-OH compounds. Both phosphatidyl-inosose and inosose types may have temporary protecting groups other than benzyls so as to avoid metal catalyzed hydrogenolysis for deprotection and concomitant reduction of C--C unsaturation in the fattyacyl chains. The present invention discloses novel selectively protected chiral myoinositol synthons that incorporate temporary protecting groups which are removed without metal catalyzed hydrogenation. In addition, the groups are compatible with the reagents and conditions validated in Schemes 1 and 2 for the oxidation and reduction steps.
- DETD Similarly, the 2-phosphatidyl isomer 1D-2-(1',2'-0-dipalmitoyl-sn-glycero-3'-phospho)-3,6di-0-benzyl-myo-inositol
 -4,5-bis(dibenzylphosphate) 51 was esterified to 52. Both 49 and 52 formed on oxidation the corresponding inosose derivative, 50 and 53 respectively. These phosphatidyl-inosose esters represent another group of key intermediates for labeling with hydrogen isotopes.
- Oxidation of 1D-1-(1',2'-O-dipalmitoyl-sn-glycero-3'-phospho)3,6-di-O-benzyl-myo-inositol-4,5bis(dibenzylphosphate) 6 by CrO.sub.3.Py.sub.2 (Procedure A) at r.t. for
 5 min was quenched by ice cold aqueous SO.sub.2. The product recovered
 by evaporation of the organic layer. Purification by chromatography on
 flash silica using a gradient of CHCl.sub.3--MeOH--NH.sub.4OH gave
 1D-1-(1', 2'-O-dipalmitoyl-sn-glycero-3'-phospho)-3,6-di-O-benzyl-2-myoinosose-4,5-bis(dibenzylphosphate) 7 (yield 69%).
- inosose-4,5-bis(dibenzylphosphate) 7 (yield 69%).

 DETD Oxidation of 1D-2-(1',2'-O-dipalmitoyl-sn-glycero-3'-phospho)3,6-di-myo-inositol-4,5-bis(dibenzylphosphate) 51 by
 Procedure A was complete at r.t. in min; worked up and purification as
 in Example 1, gave 1D-2-(1',2'-O-dipalmitoyl-sn-glycero-3'-phospho)-3,6di-1-myo-inosose-4,5-bis(dibenzylphosphate) (yield 86%).
- DETD Oxidation of 1D-1-(1',2'-0-dipalmitoyl-sn-glycero-3'-phospho)3,6-di-0-benzyl-myo-inositol-4,5bis(dibenzylphosphate)-benzyl ester 49 using Procedure A, and
 purification as in the general protocol gave 1D-1-(1',2'-0-dipalmitoylsn-glycero-3'-phospho)-3,6-di-0-benzyl-2myo-inosose
 -4,5-bis(dibenzylphosphate)-benzyl ester 50 (yield 65%).

Oxidation of 1D-2-(1',2'-O-dipalmitoyl-sn-glycero-3'-phospho)-DETD 3.6-di-O-benzyl-myo-inositol-4,5bis(dibenzy)phosphate)-benzyl ester 52 by Procedure A, work up and purification as in the general protocol gave 1D-2-(1',2'-O-dipalmitoylsn-glycero-3'-phospho)-3,6-di-0-benzyl-1-myo-inosose -4.5-bis(dibenzylphosphate)-benzyl ester 53 (yield 72%).

L84 ANSWER 4 OF 10 USPATFULL on STN

ACCESSION NUMBER:

2000:174129 USPATFULL

TITLE:

Preparation for the application of agents in

....

DATE

mini-droplets

INVENTOR(S):

Cevc, Gregor, Heimstetten, Germany, Federal Republic of

PATENT ASSIGNEE(S):

Idea AG, Munich, Germany, Federal Republic of (non-U.S.

corporation)

......

	NOWREK	KTND	DATE	
PATENT INFORMATION:	US 6165500		20001226	
APPLICATION INFO.:	US 1992-844664		19920408	(7)

		NOMBER	DATE
PRIORITY	INFORMATION:	DE 1990-4026834	19900824
		DE 1990-4026833	19900824
		DE 1991-4107153	19910306
		WO 1991-EP1596	19910822
	T-(DC	11 m 2 T 2 m m m	

DOCUMENT TYPE: FILE SEGMENT:

Utility Granted

PRIMARY EXAMINER:

Kishore, Gollamudi S.

MIMDED

LEGAL REPRESENTATIVE:

Davidson, Davidson & Kappel, LLC

NUMBER OF CLAIMS:

35

EXEMPLARY CLAIM: NUMBER OF DRAWINGS:

31 Drawing Figure(s); 21 Drawing Page(s)

LINE COUNT: 4336

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The invention relates to a preparation for the application of agents in the form of minuscule droplets of fluid, in particular provided with membrane-like structures consisting of one or several layers of amphiphilic molecules, or an amphiphilic carrier substance, in particular for transporting the agent into and through natural barriers such as skin and similar materials. The preparation contains a concentration of edge active substances which amounts to up to 99 mol-% of the agent concentration which is required for the induction of droplet solubilization. Such preparations are suitable, for example, for the non-invasive applications of antidiabetics, in particular of insulin. The invention, moreover, relates to the methods for the preparation of such formulations.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

- . K. (1989) Arzneim. Forsch./Drug Res. 39, 1487-1491). In the SUMM case of plants, improved penetration into or through the cuticle could reduce the drug concentration required for a given application and thus significantly diminish pollution problems (Price, C. E. (1981) In: The.
- form of a cyclic lactone residue. The aldehyde- or keto-groups DETD in a derivatised mono- or disaccharide residue can also be reduced to a hydroxy group, e.g. in inositol, sorbitol or D-mannitol; also, one or several hydroxy groups can be replaced by. .
- the form of cyclic lactone residues. The aldehyde- or DETD keto-groups in a derivatised mono- or disaccharide residue, moreover, can be reduced to hydroxy groups, e.g. in inositol, sorbitol or D-mannitol. Furthermore, individual hydroxy groups can be replaced by hydrogen atoms, e.g..

A carbohydrate can result from a cleaving action, starting with one of DETD the mentioned mono- or disaccharides, by a strong oxidation agent, such as periodic acid. Amongst the biologically most important or most active carbohydrates are e.g. 2-acetamido-N-(epsilon-amino-caproyl)-2-deoxy-beta-gluccopyranosylamine, 2-acetamido-2-amino-1,2-dideoxy-betaglucopyranose, 2-acetamido-1-beta-(aspartamido)-1,2-dideoxyglucose, 2-acetamido-4,6-o-benzyliden-2-deoxybeta-glucopyranose,... beta-glucopyranoside, hesperidin, n-hexyl-beta-glucopyranoside, hyaluronic acid, 16-alpha-hydroxyestronglucuronide, 16-betahydroxyestron glucuronide, hydroxyethyl starch, hydroxypropylmethylcellulose, 8-hydroxyquinolin-beta-glucopyranoside, 8-hydroxyquinolin glucuronide, idose, (-)-idose, indole-3- lactic acid, indoxyl-beta-glucoside, epi-inositol, myoinositol, myo-inositol bisphosphate, myo-inositol-1,2-cyl phosphate, scyllo-inositol, inositolhexaphosphate, inositolhexasulfate, myo-insoitol 2-monophosphate, myo-inositol trisphosphate, (q)-epi-inosose-2, scyllo-inosose, inulin, isomaltose, isomaltotriose, isosorbid dinitrate, 11-ketoandrosterone beta-glucuronide, 2-ketogluconic acid, 5-ketogluconic acid, alpha-ketopropionic acid, lactal, lactic acid, lactitol, lactobionic acid, lacto-N-tetraose,. . . acid, neuraminic acid beta-methylglycoside, neuramine-lactose, nigeran, nigerantetrasaccharide, nigerose, n-nonyl glucoside, n-nonylbeta-glucopyranoside, octadecylthio-ethyl 4-o-alphagalactopyranosyl-beta-galactopyranoside, octadecylthioethyl 4-o-(4-o-[6-o-alpha-glucopyranosyl-alpha-glucopyranosyl]-alphaglucopyranosyl)-beta-glucopyranoside, octanoyl n-methylglucamide, n-octyl alpha-glucopyranoside, n-octyl-beta-glucopyranoside, oxidised starch, pachyman, palatinose, panose, pentaerythritol, pentaerythritol diformal, 1,2,3,4,5-pentahydroxy, capronic acid, pentosanpolysulfate, perseitol, phenolphthalein glucuronic acid, phenolphthalein mono-beta-glucosiduron phenyl 2-acetamido-2-deoxy-alphagalactopyranoside, phenyl2-acetamido-2-deoxy-alpha-glucopyranoside,.

DETD Oxidoreductases, such as: alcohol dehydrogenase (1.1.1.1), alcohol dehydrogenase (NADP dependent) (1.1.1.2), glycerol dehydrogenase (1.1.1.6), glycerophosphate dehydrogenase (1.1.1.8), xylulose reductase (1.1.1.10), polyol dehydrogenase (1.1.1.14), sorbitol dehydrogenase (1.1.1.14), myo-inositol dehydrogenase (1.1.1.18), uridine 5'-diphosphoglucose dehydrogenase (1.1.1.22), glyoxalate reductase (1.1.1.26), lactate dehydrogenase (1.1.1.27), lactate dehydrogenase (1.1.1.28), glycerate dehydrogenase (1.1.1.29), beta-hydroxybutyrate dehydrogenase (1.1.1.30), beta-hydroxyacyl CoA dehydrogenase (1.1.1.35), malate dehydrogenase (1.1.1.37), glutamic dehydrogenase (1.4.1.3), glutamate dehydrogenase (NADP) (1.4.1.4), L-amino acid oxidase (1.4.3.2), D-amino acid oxidase (1.4.3.3), monoaminoxidase (1.4.3.4), diaminoxidase (1.4.3.6), dihydrofolate reductase (1.5.1.3), 5,10methylenetetrahydrofolat dehydrogenase (1.5.1.5), saccharopine dehydrogenase NAD+ (1.5.1.7), octopine dehydrogenase (1.5.1.11), sarcosine oxidase (1.5.3.1), sarcosine dehydrogenase (1.5.99.1), glutathione reductase (1.6.4.2), ferridoxin-NADP+ reductase (1.6.7.1), NADPH-FMN oxidoreductase (1.6.99.1), cytochrome c reductase (1.6.99.3), NADH-fmn oxidoreductase (1.6.99.3), dihydropteridin reductase (1.6.99.7), uricase (1.7.3.3), diaphorase (1.8.1.4), lipoamide dehydrogenase (1.8.1.4), cytochrome oxidase (1.9.3.1), nitrate reductase (1.9.6.1), phenolase (1.10.3.1), ceruloplasmine (1.10.3.2), ascorbate oxidase (1.10.3.3), NADH peroxidase (1.11.1.1), catalase (1.11.1.6), lactoperoxidase (1.11.1.7), myeloperoxidase (1.11.1.7), peroxidase (1.11.1.7), glutathione. . . salicylate hydroxylase (1.14.13.7), p-hydroxybenzoate hydroxylase (1.14.13.2), luciferase (bacterial) (1.14.14.3), phenylalanine hydroxylase (1.14.16.1), dopamine-betahydroxylase (1.14.17.1), tyrosinase (1.14.18.1), superoxide dismutase (1.15.1.1), ferredoxine-NADP reductase (1.18.1.2), etc... Transferases, such as: catecholic o-methyltransferase (2.1.1.6), phenylethanol-amine N-methyl-transferase (2.1.1.28), aspartate transcarbamylase (2.1.3.2), ornithine carbamyltransferase (2.1.3.3), transketolase (2.2.1.1), transaldolase.

. cobra), Naja Naja kaouthia, Mycoplasma gallisepticum, Perseau DETD americana (avocado). Phaseolus coccineus (beans). Phaseolus limensis. Phaseolus lunatus, Phaseolus vulgaris, Phytolacga americana, Pseudomonas aeruginosa PA-I, Pisum sativum (pea), Ptilota plumosa (red algae), Psophocarpus tetragonolobus (winged bean), Ricinus communis (castor bean), Robinia pseudoacacia (false.

. . be parts of a biological extract. As sources of biologically DETD and/or pharmacologically active extracts, the following are worth-mentioning: for example, Acetobacter pasteurianum, Acokanthera ouabaio cathel, Aesculus hippocastanum, Ammi visnaga Lam., Ampi Huasca, Apocynum Cannabium, Arthrobotrys superba var. oligospora (ATCC 11572), Atropa.

Next, the carrier composition or concentration is adapted by DETD reducing the edge activity in the system to an extent which ensures the vesicle stability as well vesicle deformability to be. .

If the pore diameter is reduced to 0.05 micrometers only suspensions with L/S ratios below 2/1 can still be filtered.

L84 ANSWER 5 OF 10 USPATFULL on STN

ACCESSION NUMBER:

2000:40863 USPATFULL

TITLE:

Highly sensitive method for assaying chiro-inositol and

compositions for the assay

INVENTOR(S):

Kozuma, Takuji, Shizuoka, Japan Takahashi, Mamoru, Shizuoka, Japan

PATENT ASSIGNEE(S):

Asahi Kasei Kogyo Kabushiki Kaisha, Osaka, Japan

(non-U.S. corporation)

	NUMBER	KIND	DATE	
PATENT INFORMATION:	US 6046018		20000404	
	WO 9842863		19980110	
APPLICATION INFO.:	US 1999-308575		19990608	(9)
	WO 1998-JP1215		19980320	
			19990608	PCT 371 date
			19990608	PCT 102(e) da

9990608 PCT 102(e) date

NUMBER DATE

PRIORITY INFORMATION:

JP 1997-72878 19970326

DOCUMENT TYPE:

Utility Granted

FILE SEGMENT:

PRIMARY EXAMINER: LEGAL REPRESENTATIVE:

Leary, Louise N. Young & Thompson

NUMBER OF CLAIMS:

5

EXEMPLARY CLAIM:

1

NUMBER OF DRAWINGS:

6 Drawing Figure(s); 6 Drawing Page(s)

LINE COUNT: 1012

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The present invention relates to an assay method of chiroinositol which AB comprises reacting a specimen containing chiroinositol with

- 1) a dehydrogenase, which catalyses at least reversible reaction with a substrate of chiroinositol in the presence of a coenzyme selected from NAD(P)s and a coenzyme selected from thio-NAD(P)s,
- 2) A1 and
- 3) B1

to form cycling reaction of the formula ##STR1## wherein a product is a compound, from which 2 or 4 hydrogen atoms are deleted from chiroinositol, A1 is NAD(P)s or thio-NAD(P)s, A2 is a reduced form of A1, B1 is a reduced form of NAD(P)s in case of A1 being thio-NAD(P)s or a reduced form of thio-NAD(P)s in case of A1 being NAD(P)s and B2 is an oxidized form of B1, and determining an amount of converted A2 or B1 by the said reaction, and a composition for assay of chiroinositol. Chiroinositol can be assayed by accurate,

simple, low price and high sensitive method.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

```
. compound, from which 2 or 4 hydrogen atoms are deleted from
AB
       chiroinositol, A1 is NAD(P)s or thio-NAD(P)s, A2 is a reduced
       form of A1, B1 is a reduced form of NAD(P)s in case of A1
       being thio-NAD(P)s or a reduced form of thio-NAD(P)s in case
       of A1 being NAD(P)s and B2 is an oxidized form of B1, and determining
             . compound, from which 2 or 4 hydrogen atoms are deleted from
SUMM
       chiroinositol, A1 is NAD(P)s or thio-NAD(P)s, A2 is a reduced
       form of A1, B1 is a reduced form of NAD(P)s in case of A1
       being thio-NAD(P)s or a reduced form of thio-NAD(P)s in case
       of Al being NAD(P)s and B2 is an oxidized form of B1, and determining
SUMM
       3) in the above 2), at least a coenzyme selected from reduced
       thio-NAD (P)s in case of at least a coenzyme selected from NAD(P)s, or
       in the above 2), at least a coenzyme selected from reduced
       NAD(P)s in case of at least ,t coenzyme selected from thio-NAD(P)s.
SUMM
             in body fluid such as blood or urine for diagonosis of diabetes
       mellitus, especially insulin resistance, is useful, and suggest
       reduction/oxidation analysis using enzyme, however no concrete
       method has proposed.
SUMM
       In order to assay trace amount of chiroinositol in vivo in clinical
       biochemical test, not only direct assay method of reduced
       coenzyme using dehydrogenase but also a combination with coloring agent
       for assay is resulted to insufficient sensitivity. We have found.
       dehydrogenase derived from Aerobacter aerogenes acts on myoinositol in
       the presence of NAD to delete 2 hydrogen atoms to form
       myoinosose 2, under sufficient progressive condition for
       reaction, a compound generated from a reaction in which at first 2
       hydrogen atoms. . . not detected by paper chromatography [J. Biol. Chem., 241 (4); 1966, 800-806]. Since the said final compound is
       different from myoinosose 2 and is very unstable, to construct
       a stable enzymatic cycling reaction might be impossible.
SUMM
            . compound, from which 2 or 4 hydrogen atoms are deleted from
       chiroinisitol, A1 is NAD(P)s or thio-NAD(P)s, A2 is a reduced
       form of A1, B1 is a reduced form of NAD(P)s in case of A1
       being thio-NAD(P)s or a reduced form of thio-NAD(P)s in case
       of A1 being NAD(P)s and B2 is an oxidized form of B1, and can be.
SUMM
             . compound, from which 2 or 4 hydrogen atoms are deleted from
       chiroinositol, A1 is NAD(P)s or thio-NAD(P)s, A2 is a reduced
       form of A1, B1 is a reduced form of NAD(P)s in case of A1
       being thio-NAD (P)s or a reduced form of thio-NAD(P)s in case
       of A1 being NAD(P)s and B2 is an oxidized form of B1, and determining
SUMM
       3) in the above 2), at least a coenzyme selected from reduced
       thio-NAD(P)s in case of at least a coenzyme selected from NAD(P)s, or in
       the above 2), at least a coenzyme selected from reduced
       NAD(P)s in case of at least a coenzyme selected from thio-NAD(P)s.
        . . . production (detected by lead acetate paper) -
SUMM
  Acetoin production (K.sub.2 HPO.sub.4) -
  Acetoin Production (NaCl) -
  MR test -
  Nitrate reduction test (gas formation) -
  (NO.sub.2 - detection)
  (NO.sub.3 - detection) +
  Utilization on Simmons medium
  Citrate -
  Malate -
              C. + + +
  55.degree. C. + + +
  60.degree. C. ND ND -
  70.degree. C. - - +
  Nitrate reduction d + -
  GC mole % of DNA 44.5 46.4 41.9
```

(Type) (Type) 44.3.about.50.3 42.9.about.49.9

SUMM The present enzyme catalyses a reaction for generating **reduced** coenzyme [NAD(P)Hs and thio-NAD(P)Hs] in the presence of chiroinositol and coenzyme [NAD(P)s and thio-NAD(P)s]. Examples of the above NAD(P)s are. . .

SUMM . . . and deamino NAD; 14 U/ml) 20 .mu.l is added and stirred.
Absorption changes per minute in specific wavelength for each
reduced coenzyme is measured to obtain initial reaction rate.
(For NAD and deamino NAD, measured value is increased number by ten.

SUMM A product in the present cycling reaction is an amount of reduced NAD generated by the reaction with chiroinositol and excess amount of NAD. In the reaction, 2 hydrogen atoms are deleted.

. the first reaction and 2 hydrogen atoms are further deleted at the second reaction. These are confirmed by increase in reduced NAD, which is determined by an amount of formazan pigment having maximum absorption at 550 nm generated as a result of an act on of NBT (nitroblue tetrazolium) on the reduced NAD in the presence of diaphorase.

SUMM

TABLE 1

Substrate Relative activity

chiroinositol 100%
myoinositol 9%
scylloinositol 0%
epi-inositol 0%
galactose 0%
fructose 0%
mannose 0%
mannitol 0%

SUMM

TABLE 3

A.r.1215 origin S.r.301 origin

100%

chiroinositol 100%
myoinositol 33% 0%
scylloinositol 0% 0%
epi-inositol 0% 4%
fructose 0% 0%
mannose 10% 0%
mannitol 0% 0%

SUMM . . . 2 or 4 hydrogen atoms are deleted from chiroinositol, A1 and B2 are NAD(P)s or thio-NAD(P)s, A1 and B1 are reduced form thereof, and in A1 and B1, when A1 is thio-NAD(P)s, B1 is NAD(P)Hs, and when B1 is thio-NAD(P)H, A1. . .

SUMM In case of enzyme cycling method in the present invention, if A1 and B1 are expensive, in order to reduce amount of A1 and B1, a combination of a dehydrogenase which constitutes a reaction of B2.fwdarw.B1 and not reacted with. . . dehydrogenase which constitutes a reaction of A2.fwdarw.A1 and not reacted with chiroinositol and substrate for dehydrogenase can be used for reducing amount of A1 and B1.

SUMM Reduced coenzyme assay by measuring absorption change used in the present invention can be performed by other known method.

DETD As shown in the above, the present invention provide rate assay of the reduced coenzyme and the blank assay for the specimen can be ommitted. Consequently, simple assay can be performed, and sensitivity of. . .

CLM What is claimed is:

. compound, from which 2 or 4 hydrogen atoms are deleted from chiroinositol, A1 is NAD(P)s or thio-NAD(P)s, A2 is a reduced

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form of A1, B1 is a reduced form of NAD(P)s in case of A1 being thio-NAD(P)s or a reduced form of thio-NAD(P)s in case of Al being NAD(P)s and B2 is an oxidized form of B1, and determining

. least a coenzyme selected from NAD(P)s and thio-NAD(P)s, and 3) in the above 2), at least a coenzyme selected from reduced thio-NAD(P)s in case of at least a coenzyme selected from NAD(P)s, or in the above 2), at least a coenzyme selected from reduced NAD(P)s in case of at least a coenzyme selected from thio-NAD(P)s.

L84 ANSWER 6 OF 10 USPATFULL on STN

ACCESSION NUMBER:

94:90948 USPATFULL

TITLE:

Highly sensitive assay method for myo-inositol, composition for practicing same, novel myo-inositol

dehydrogenase, and process for producing same Ueda, Shigeru, Shizuoka, Japan

INVENTOR(S):

Takahashi, Mamoru, Shizuoka, Japan Misaki, Hideo, Shizuoka, Japan Imamura, Shigeyuki, Shizuoka, Japan Matsuura, Kazuo, Shizuoka, Japan

PATENT ASSIGNEE(S):

Asahi Kasei Kogyo Kabushiki Kaisha, Osaka, Japan

(non-U.S. corporation)

NUMBER KIND DATE

PATENT INFORMATION:

US 5356790 US 1993-106693

19941018 19930816 (8)

APPLICATION INFO.: RELATED APPLN. INFO.:

Continuation of Ser. No. US 1991-761465, filed on 18

Sep 1991, now abandoned

NUMBER

PRIORITY INFORMATION:

JP 1990-2249775 19900918 19900918 JP 1990-2249776

Utility

DOCUMENT TYPE: FILE SEGMENT:

Granted

PRIMARY EXAMINER: ASSISTANT EXAMINER:

Wityshyn, Michael G. Leary, Louise N. Young & Thompson

LEGAL REPRESENTATIVE: NUMBER OF CLAIMS:

EXEMPLARY CLAIM:

NUMBER OF DRAWINGS:

8 Drawing Figure(s); 8 Drawing Page(s)

LINE COUNT: 819

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Myo-inositol in a specimen is assayed by reacting a specimen containing AB mvo-inositol with:

- a) myo-inositol dehydrogenase using a thio-NADP group or thio-NAD group and an NADP group or NAD group as coenzymes, and which catalyzes a reversible reaction forming myo-inosose from myo-inositol.
- b) A.sub.1 and
- c) B.sub.1

to effect a cycling reaction ##STR1## wherein A.sub.1 is a thio-NADP group, thio-NAD group, NADP group or NAD group, A.sub.2 is a reduced form of A.sub.1, when A.sub.1 is a thio-NADP group or thio-NAD group, B.sub.1 is a reduced NADP group or reduced NAD group and when A.sub.1 is an NADP group or NAD group, B.sub.1 is a reduced thio-NADP group or reduced thio-NAD group, and wherein B.sub.2 is an oxidized form of B.sub.1. The change in the amount of A.sub.2 generated or B.sub.1 consumed by the cycling reaction is measured to perform the assay. A composition for performing the assay comprises the above myo-inositol dehydrogenase, as well as the above components A.sub.1 and B.sub.1. The

myo-inositol dehydrogenase can be produced by culturing a suitable microorganism belonging to genus Bacillus, particularly Bacillus sp. No. 3 FERM BP-3013.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

- AB a) myo-inositol dehydrogenase using a thio-NADP group or thio-NAD group and an NADP group or NAD group as coenzymes, and which catalyzes a reversible reaction forming myo-inosose from myo-inositol.
- SUMM (1) myo-inositol dehydrogenase using one of coenzymes of thionicotinamide adonine dinucleotide phosphate group (hereinafter designated thio-NADP group) or thionicotinamide adenine dinucleotide group. . . (hereinafter designated NADP group) or nicotinamide adonine dinucleotide group (hereinafter designated NAD group) and which catalyzes a reversible reaction forming myo-inosose from a substrate of myo-inositol,
- DETD Aerobacter aerogenes (J. Biol. Chem., 241, 800-806 (1966)); Klebsiella pneumoniae, Serratia marcescens, Cryptococcus melibiosum (Biochim. Biophys. Acta., 293, 295-303 (1973)); and bovine brain (Biochem. Biophys. Res. Comm., 68, 1133-1138 (1976)); Bacillus. . .
- DETD Among these, Aerobacter aerogenes, Klebsiella pneumoniae and Serratia marcescens are known as etiologic microorganisms for pneumonia and opportunistic infections (Standard Microbiology, 2nd edn., pp. 209-212, Igaku Shoin Publishing. . .
- DETD The enzyme catalyzes essentially a reaction of myoinositol and NAD to generate myo-inosose and reduced NADH. as follows:
- DETD myo-inositol+NAD.about.myo-inosose
 *+reduced NADH *(2,4,6/3,5-pentahydroxy cyclohexanone)
- DETD Glyoxylate dehydrogenase (EC.1.2.1.17) (Pseudomonas oxalaticus) and CoA and glyoxylate,
- DETD Benzaldehyde dehydrogenase (EC.1.2.1.7) (Pseudomonas fluorescens) and benzaldehyde.
- CLM What is claimed is: 1. A method of assaying myo-inositol comprising reacting a specimen containing myo-inositol with the following reagents: a) myo-inositol dehydrogenase which, in the presence of a thionicotinamide adenine dinucleotide group (thio-NAD-group) and a nicotinamide adenine dinucleotide group (NAD group) as coenzymes, catalyzes a reversible reaction forming myoinosose from myo-inositol, b) A.sub.1 and c) B.sub.1 ; to effect a cycling reaction ##STR5## wherein A.sub.1 is a thio-NAD group or NAD. . . NAD group and when A.sub.1 is an NAD group, B.sub.1 is a reduced thio-NAD group, and wherein B.sub.2 is an oxidized form of B.sub.1; and measuring a change in the amount of A.sub.2 generated or B.sub.1 consumed by the cycling reaction wherein A.sub.1 and B.sub.1 are each used at a concentration of 0.02-100 mM, and wherein said myo-inositol dehydrogenase is used at a concentration of 5-1000 U/ml.
 - 3. A reagent composition for assaying myo-inositol, comprising: a) myo-inositol dehydrogenase which, in the presence of a thionicotinamide adenine dinucleotide group (thio-NAD group) and a nicotinamide adenine dinucleotide group (NAD group) as coenzymes, catalyzes a reversible reaction forming myo-inosose from myo-inositol, b) A.sub.1 and c) B.sub.1; wherein A.sub.1 is a thio-NAD group or NAD group, when A.sub.1 is a thio-NAD. . . of a thio-NAD group wherein A.sub.1 and B.sub.1 are each present in a concentration of 0.02-100 mM, and wherein said myo-inositol dehydrogenase is present in a concentration of 5-1000 U/ml.
 - 4. Myo-inositol dehydrogenase having the following properties: substrate spcificity for myo-inositol and catalyzes a reaction myo-inositol+NAD.about.myo-inosose+reduced NADH, said myo-inositol dehydrogenase having the following physicochemical properties: (1)

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molecular weight: 130,000.+-.15,000 (gel filtration method by TSK gel G 3000 SW) (2) iso-electric point: pH 4.5.+-.0.5 (3) Km-value: Km value for myo-inositol: 0.64 mM Km value for NAD: 0.004 mM (4) optimum pH: approximately ph. 9.5 (5) pH-stability: more than 80% retained. .

L84 ANSWER 7 OF 10 USPATFULL on STN

ACCESSION NUMBER:

93:57049 USPATFULL

TITLE: INVENTOR(S): 3-deoxy-3-substituted analogs of phosphatidylinositol Kozikowski, Alan P., Ponte Verde Beach, FL, United

Tuckmantel, Werner, Jacksonville, FL, United States Faug, Abdul H., Jacksonville, FL, United States

Powis, Garth, Tucson, AZ, United States

PATENT ASSIGNEE(S):

Mayo Foundation for Medical Education and Research.

Rochester, MN, United States (U.S. corporation)

NUMBER KIND DATE -----19930713

PATENT INFORMATION: APPLICATION INFO.:

US 5227508 US 1992-825523 19920124 (7)

DOCUMENT TYPE: Utility FILE SEGMENT: Granted

Ramsuer, Robert W. PRIMARY EXAMINER: ASSISTANT EXAMINER: Ambrose, Michael G.

LEGAL REPRESENTATIVE:

Merchant, Gould, Smith, Edell, Welter & Schmidt 15

NUMBER OF CLAIMS: EXEMPLARY CLAIM: 1

NUMBER OF DRAWINGS: 6 Drawing Figure(s): 6 Drawing Page(s)

927 LINE COUNT:

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The invention provides 3-deoxy-3-substituted analogs of

phosphatidylinositol which are useful to inhibit the growth of mammalian cells, i.e., to treat neoplastic conditions and other proliferative disorders of mammalian cells.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

The key intermediate, 2,4,5,6-tetra-O-benzyl-3-deoxy-3-fluoro-Dmyo-inositol (40), is available as outlined earlier. Inversion of the stereochemistry at C-1 is brought about by oxidation to the inosose ((COC1).sub.2, DMSO,

i-Pr.sub.2 NEt), followed by stereoselective reduction of the 1-ketone with L-Selectride.RTM. (Aldrich Chem. Co.). The resulting axial alcohol.

L84 ANSWER 8 OF 10 USPATFULL on STN

ACCESSION NUMBER:

76:52995 USPATFULL

TITLE:

Process for preparing aminocyclitol antibiotics

INVENTOR(S):

Daum, Sol J., Albany, NY, United States

Clarke, Robert L., Bethlehem, NY, United States Sterling Drug Inc., New York, NY, United States (U.S.

PATENT ASSIGNEE(S):

corporation)

NUMBER KIND DATE ______ PATENT INFORMATION: US 3982996 19760928 APPLICATION INFO.: US 1975-615593 19750922 (5)

DOCUMENT TYPE: FILE SEGMENT:

Utility Granted

PRIMARY EXAMINER:

Tanenholtz, Alvin E.

LEGAL REPRESENTATIVE:

Webb, William G., Wyatt, B. W.

NUMBER OF CLAIMS: EXEMPLARY CLAIM:

.12 1

LINE COUNT:

728

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

Aminocyclitol antibiotics of the streptamine, deoxystreptamine or dideoxystreptamine type are prepared by culturing a nutrient medium

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containing carbohydrates, a source of assimilable nitrogen, essential salts and a non-nitrogen containing cyclitol with a mutant of an aminocyclitol antibiotic producing organism.

CAS INDEXING IS AVAILABLE FOR THIS PATENT.

SUMM Yet a process that would permit the use of cyclitols, instead of aminocyclitols, for incorporation into aminocyclitol-type antibiotics by microorganism mutants using the Rinehart/Shier method would provide a very significant advance in the aminocyclitol antibiotic art, because the method would afford, by judicious selection of the microorganism and the cyclitol subunit, a certain degree of biogenetic "tailoring" of the resultant antibiotic molecule. Moreover, since the aminocyclitols are. . . products could be realized. (For example, streptamine, at present prices, costs about \$1 per gram, whereas its probable biogenetic precursor, scyllo-inosose, can be obtained in about 80% yield by fermentative oxydation of myo-inositol, which only costs about 2 cents per gram at present).

DETD . . . [dl-1,2,3,4,5-cyclohexanepentol (1,2,4-cis)] [McCasland et al., J. Am. Chem. Soc. 75, 4020 (1953)] (0.40 mole) was subjected to microbiological oxidation by Acetobacter suboxydans using the procedure described by Posternak, Helv. Chim. Acta 33, 1594-1596 (1950). To the resulting broth was added 5. . .

DETD The latter was subjected to microbiological oxidation by

Acetobacter suboxydans using the procedure described by

Posternak recorded above in Preparation 1, and the product was isolated as described in. . .

=> d ibib abs 9

L84 ANSWER 9 OF 10 SCISEARCH COPYRIGHT 2003 THOMSON ISI on STN

ACCESSION NUMBER: 97:313486 SCISEARCH

THE GENUINE ARTICLE: WT993

TITLE: Reactions of the ketone derived from (+/-)-3,4,5-tri-0-

benzyl-1,2-0-isopropylidene-myo-inositol: Preparation of

racemic derivatives of epi-inositol

and of 4-C-methyl-epi-[(+/-)-iso-laminitol] and

4-C-methyl-myo-inositol [(+/-)-laminitol]

AUTHOR: Gigg J; Gigg R (Reprint)

CORPORATE SOURCE: NATL INST MED RES, DIV LIPID & GEN CHEM, MILL HILL, LONDON

NW7 1AA, ENGLAND (Reprint); NATL INST MED RES, DIV LIPID &

GEN CHEM, LONDON NW7 1AA, ENGLAND

COUNTRY OF AUTHOR: ENGLAND

SOURCE: CARBOHYDRATE RESEARCH, (26 MAR 1997) Vol. 299, No. 1-2,

pp. 77-83.

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LANGUAGE:

English

REFERENCE COUNT: 27

ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS

AB Oxidation of (+/-)-3,4,5-tri-0-benzyl-1,2-0-isopropylidene-myo-inositol with the pyridine-SO3 complex in methyl sulfoxide gave the ketone which was reduced with sodium borohydride to give almost exclusively the corresponding epi-inositol derivative. Reaction of the ketone with diazomethane gave an epoxide which was reduced with lithium aluminium hydride to give a 4-C-methyl-myo-inositol derivative and reaction of the ketone with methyl magnesium iodide gave the isomeric 4-C-methyl-epi-inositol derivative. (C) 1997 Elsevier Science Ltd.

L84 ANSWER 10 OF 10 WPIX COPYRIGHT 2003 THOMSON DERWENT on STN

ACCESSION NUMBER:

2002-158901 [21] WPIX

DOC. NO. CPI:

C2002-049937

TITLE:

L-epi-Inositol derivative is useful as an intermediate of medicaments or agrochemicals.

DERWENT CLASS: PATENT ASSIGNEE(S): B05 C03 (HOKK) HOKKO CHEM IND CO LTD

COUNTRY COUNT:

PATENT INFORMATION:

PG PATENT NO KIND DATE WEEK LA JP 2001335544 A 20011204 (200221)*

APPLICATION DETAILS:

PATENT NO APPLICATION DATE KTND JP 2001335544 A JP 2000-158238 20000529

PRIORITY APPLN. INFO: JP 2000-158238 20000529

2002-158901 [21] WPIX

AB JP2001335544 A UPAB: 20020403

> NOVELTY - A L-epi-inositol derivative or its salt (I), are new.

DETAILED DESCRIPTION - A L-epi-inositol derivative of formula (I) or its salt, are new.

R1, R4-R7 = H, acyl or alkyl;

when R2 = amino, acylamide, alkylamino or N-acyl-N-alkylamino, R3 H; when R2 = hydroxyl or acyloxyl, R3 = hydroxymethyl, acyloxymethyl, azidemethyl, aminomethyl, acylamidemethyl, N-alkylaminomethyl or N-acyl-N-alkylaminomethyl; when R2 = 0, R3 methylene, R3 + R2 +spirocarbon at the second position of the cyclohexane ring bind to each other to form 2,21-anhydro-2-C-hydroxymethyl of spiroepoxy ring.

INDEPENDENT CLAIMS are also included for a method of preparing 2-amino-2-deoxy-L-epi-inositol of formula (II) which comprises allowing the ketone group of L-epi-2inosose of formula (III) to react with an ammonia derivative for dehydration condensation, and reducing with a reducing agent in the presence of a catalyst to convert into an amino group.

USE - The inositol derivative is useful as an intermediate of medicaments or agrochemicals.

ADVANTAGE - The inositol derivative having biological activity is inexpensively prepared in high yields.

Dwg.0/0

MARX 09/980,453

=> d his (FILE 'HOME' ENTERED AT 14:25:34 ON 18 AUG 2003) FILE 'CAPLUS' ENTERED AT 14:26:20 ON 18 AUG 2003 . L1 65 S KANBE K?/AU L2 4489 S TAKAHASHI A?/AU L3 8664 S MORI T?/AU 457 S TAMAMURA T?/AU L4 L5 6978 S TAKEUCHI T?/AU L6 1234 S KITA Y?/AU L7 21776 S L1-6 L8 3 S L7 AND EPI-INOSITOL 7 S L7 AND ?INOSOSE 19 7 S L8-9 L10 SELECT RN L10 1-7 FILE 'REGISTRY' ENTERED AT 14:29:26 ON 18 AUG 2003 L11 36 S E1-36 FILE 'CAPLUS' ENTERED AT 14:29:32 ON 18 AUG 2003 36 cpds displayed L12 7 S L10 AND L11 7 cites 4 FILE 'CAPLUS' ENTERED AT 14:29:54 ON 18 AUG 2003 => d ibib abs hitstr ind 1-7 L12 ANSWER 1 OF 7 CAPLUS COPYRIGHT 2003 ACS on STN ACCESSION NUMBER: 2003:266870 CAPLUS DOCUMENT NUMBER: 138:270409 TITLE: Scyllo-inosose and scyllo-inositol manufacture INVENTOR(S): Kamibe, Kenji; Takahashi, Atsushi; Kita, Yuichi; Yamaguchi, Masanori; Tamamura, Takeshi; Mori, Tetsuya Hokko Chemical Industry Co., Ltd., Japan Jpn. Kokai Tokkyo Koho, 16 pp. PATENT ASSIGNEE(S): SOURCE: CODEN: JKXXAF DOCUMENT TYPE: Patent LANGUAGE: Japanese FAMILY ACC. NUM. COUNT: PATENT INFORMATION: PATENT NO. KIND DATE APPLICATION NO. DATE JP 2003102492 20030408 JP 2002-184912 A2 20020625 PRIORITY APPLN. INFO.: JP 2001-191161 A 20010625 The scyllo-inosose is manufd. from myo-inositol with Pseudomos and Acetobacter. The scyllo-inosose is reduced with an reductant such as sodium borohydride to get scyllo-inositol. The physiol. and morphol. characteristics of these microorganisms were given. scyllo-inosose is an useful intermediate for manufq. pharmaceuticals. The scyllo-inositol is useful for control of. Alzheimer disease and for prepd. liq. crystal. IT 488-64-2P, scyllo-Inosose RL: BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses) (scyllo-inosose and scyllo-inositol manuf.) 488-64-2 CAPLUS

Relative stereochemistry.

CN

myo-2-Inosose (7CI, 9CI) (CA INDEX NAME)

IT

87-89-8, myo-Inositol
RL: BSU (Biological study, unclassified); RCT (Reactant); BIOL (Biological study); RACT (Reactant or reagent)

(scyllo-inosose and scyllo-inositol manuf.)

87-89-8 CAPLUS RN

myo-Inositol (9CI) (CA INDEX NAME)

Relative stereochemistry.

IT 16940-66-2, Sodium borohydride

RL: RCT (Reactant); RACT (Reactant or reagent) (scyllo-inosose and scyllo-inositol manuf.)

16940-66-2 CAPLUS

Borate(1-), tetrahydro-, sodium (8CI, 9CI) (CA INDEX NAME) CN

Na+

488-59-5P, scyllo-Inositol

RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(scyllo-inosose and scyllo-inositol manuf.)

RN 488-59-5 CAPLUS

scyllo-Inositol (9CI) (CA INDEX NAME)

Relative stereochemistry.

```
ICM C12P007-26
IC
     ICS C07C029-143; C07C035-16; C12N001-20; C12P007-18; C12R001-38;
          C12R001-02
     16-2 (Fermentation and Bioindustrial Chemistry)
     Section cross-reference(s): 1
     scyllo inosose manuf myoinositol Pseudomos Acetobacter; redn
     scyllo inositol Alzheimer disease pharmaceutical
    Acetobacter
TT
     Alzheimer's disease
     Fermentation
     Liquid crystals
     Pseudomonas
     Reducing agents
        (scyllo-inosose and scyllo-inositol manuf.)
     488-64-2P, scyllo-Inosose
     RL: BPN (Biosynthetic preparation); THU (Therapeutic use); BIOL
     (Biological study); PREP (Preparation); USES (Uses)
        (scyllo-inosose and scyllo-inositol manuf.)
     87-89-8, myo-Inositol
     RL: BSU (Biological study, unclassified); RCT (Reactant); BIOL (Biological
     study); RACT (Reactant or reagent)
        (scyllo-inosose and scyllo-inositol manuf.)
     16940-66-2, Sodium borohydride
     RL: RCT (Reactant); RACT (Reactant or reagent)
        (scyllo-inosose and scyllo-inositol manuf.)
     488-59-5P, scyllo-Inositol
     RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological
     study); PREP (Preparation); USES (Uses)
        (scyllo-inosose and scyllo-inositol manuf.)
L12 ANSWER 2 OF 7 CAPLUS COPYRIGHT 2003 ACS on STN
                         2002:672204 CAPLUS
ACCESSION NUMBER:
DOCUMENT NUMBER:
                         137:200353
                         D-allo-5-inosose, its microbial manufacture,
TITLE:
                         and manufacture of allo-inositol, D-allo-3-
                         inosose, or D-chiro-inositol
                         Takahashi, Atsushi; Yamaguchi, Masanori;
INVENTOR(S):
                         Mori, Tetsuya; Kamibe, Kenji; Kita,
                         Yuichi; Tomoda, Akihiro; Tamamura,
                         Takeshi
                         Hokko Chemical Industry Co., Ltd., Japan
PATENT ASSIGNEE(S):
                         Jpn. Kokai Tokkyo Koho, 27 pp.
SOURCE:
                         CODEN: JKXXAF
DOCUMENT TYPE:
                         Patent
                         Japanese
LANGUAGE:
FAMILY ACC. NUM. COUNT:
                         1
PATENT INFORMATION:
     PATENT NO.
                      KIND
                            DATE
                                            APPLICATION NO.
                                                             DATE
     JP 2002249459
                            20020906
                                            JP 2001-46412
                                                             20010222
                       A2
PRIORITY APPLN. INFO.:
                                        JP 2001-46412
                                                             20010222
OTHER SOURCE(S):
                         CASREACT 137:200353
      OH
```

Ι

AB D-Allo-5-inosose (I), useful as a starting material for manuf.

of pharmaceuticals, is manufd. by treatment of epiinositol (II) with microorganisms capable of oxidizing II into I.
Allo-inositol (III) is manufd. by redn. of I with alkali metal
borohydrides, alkali metal trialkoxyborohydrides, or alkali metal
cyanoborohydrides as reducing agents in aq. media and sepn. of III from
II. D-Allo-3-inosose (IV) is manufd. by treatment of III with
microorganisms capable of oxidizing III into IV. D-Chiro-inositol (V),
useful for treatment of non-insulin-dependent diabetes mellitus and
polycystic ovary syndrome, is manufd. by redn. of IV with alkali metal
borohydrides, alkali metal trialkoxyborohydrides, or alkali metal
cyanoborohydrides in aq. media and sepn. of V from III. V was manufd.
from II and purified in a total yield of 32.3%.
643-10-7P, allo-Inositol

RL: BCP (Biochemical process); BMF (Bioindustrial manufacture); IMF (Industrial manufacture); PUR (Purification or recovery); RCT (Reactant); BIOL (Biological study); PREP (Preparation); PROC (Process); RACT (Reactant or reagent)

(manuf. of D-allo-5-inosose, allo-inositol, D-allo-3-inosose, and D-chiro-inositol for pharmaceuticals)

RN 643-10-7 CAPLUS

CN allo-Inositol (9CI) (CA INDEX NAME)

epi-Inositol (9CI) (CA INDEX NAME)

Relative stereochemistry.

IT 488-58-4, epi-Inositol
RL: BCP (Biochemical process); RCT (Reactant); BIOL (Biological study);
PROC (Process); RACT (Reactant or reagent)
(manuf. of D-allo-5-inosose, allo-inositol, D-allo-3inosose, and D-chiro-inositol for pharmaceuticals)
RN 488-58-4 CAPLUS

Relative stereochemistry.

IT 643-12-9P, D-chiro-Inositol
RL: BMF (Bioindustrial manufacture); IMF (Industrial manufacture); PUR
(Purification or recovery); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
(manuf. of D-allo-5-inosose, allo-inositol, D-allo-3-inosose, and D-chiro-inositol for pharmaceuticals)
RN 643-12-9 CAPLUS
CN D-chiro-Inositol (9CI) (CA INDEX NAME)

Absolute stereochemistry.

148218-11-5P, D-Allo-3-Inosose 452335-59-0P,

D-allo-5-Inosose

RL: BMF (Bioindustrial manufacture); PUR (Purification or recovery); RCT (Reactant); BIOL (Biological study); PREP (Preparation); RACT (Reactant or

(manuf. of D-allo-5-inosose, allo-inositol, D-allo-3inosose, and D-chiro-inositol for pharmaceuticals)
148218-11-5 CAPLUS

RN

D-allo-3-Inosose (9CI) (CA INDEX NAME) CN

Absolute stereochemistry.

452335-59-0 CAPLUS

D-allo-5-Inosose (9CI) (CA INDEX NAME) CN

Absolute stereochemistry.

13762-51-1, Potassium borohydride 16940-17-3, Sodium trimethoxyborohydride 16940-66-2, Sodium borohydride

16949-15-8, Lithium borohydride 25895-60-7, Sodium

cyanoborohydride

RL: RCT (Reactant); RACT (Reactant or reagent)

(reducing agent; manuf. of D-allo-5-inosose, allo-inositol, D-allo-3-inosose, and D-chiro-inositol for pharmaceuticals)

RN 13762~51-1 CAPLUS

Borate(1-), tetrahydro-, potassium (8CI, 9CI) (CA INDEX NAME)

κ+

RN 16940-17-3 CAPLUS CN Borate(1-), hydrotrimethoxy-, sodium, (T-4)- (9CI) (CA INDEX NAME)

■ Na +

RN 16940-66-2 CAPLUS CN Borate(1-), tetrahydro-, sodium (8CI, 9CI) (CA INDEX NAME)

Na⁺

RN 16949-15-8 CAPLUS CN Borate(1-), tetrahydro-, lithium (8CI, 9CI) (CA INDEX NAME)

Li+

RN 25895-60-7 CAPLUS CN Borate(1-), (cyano-.kappa.C)trihydro-, sodium, (T-4)- (9CI) (CA INDEX NAME)

Na+

```
ICM C07C049-497
     ICS C07C029-143; C07C035-16; C12P007-02; C12P007-26; C12R001-64;
          C12R001-01; C12R001-38
CC
     16-2 (Fermentation and Bioindustrial Chemistry)
     Section cross-reference(s): 63
     inosose inositol manuf fermn antidiabetic; polycystic ovary
     syndrome treatment inositol manuf; biochem oxidn redn inosose
     inositol manuf
IT
    Oxidation
        (biol.; manuf. of D-allo-5-inosose, allo-inositol, D-allo-3-
        inosose, and D-chiro-inositol for pharmaceuticals)
    Acetobacter
     Agrobacterium
     Antidiabetic agents
     Enterobacter
     Fermentation
     Gluconobacter
     Haemophilus
     Pasteurella
     Pseudomonas
     Reducing agents
     Reduction
     Serratia
     Sphingomonas
     Xanthomonas
     Yersinia
        (manuf. of D-allo-5-inosose, allo-inositol, D-allo-3-
        inosose, and D-chiro-inositol for pharmaceuticals)
TT
     Diabetes mellitus
        (non-insulin-dependent, therapeutic agents; manuf. of D-allo-5-
        inosose, allo-inositol, D-allo-3-inosose, and
        D-chiro-inositol for pharmaceuticals)
IT
    Ovary, disease
        (polycystic, therapeutic agents; manuf. of D-allo-5-inosose,
        allo-inositol, D-allo-3-inosose, and D-chiro-inositol for
        pharmaceuticals)
     643-10-7P, allo-Inositol
     RL: BCP (Biochemical process); BMF (Bioindustrial manufacture); IMF
     (Industrial manufacture); PUR (Purification or recovery); RCT (Reactant);
     BIOL (Biological study); PREP (Preparation); PROC (Process); RACT
     (Reactant or reagent)
        (manuf. of D-allo-5-inosose, allo-inositol, D-allo-3-
        inosose, and D-chiro-inositol for pharmaceuticals)
     488-58-4, epi-Inositol
     RL: BCP (Biochemical process); RCT (Reactant); BIOL (Biological study);
     PROC (Process); RACT (Reactant or reagent)
        (manuf. of D-allo-5-inosose, allo-inositol, D-allo-3-
        inosose, and D-chiro-inositol for pharmaceuticals)
     643-12-9P, D-chiro-Inositol
     RL: BMF (Bioindustrial manufacture); IMF (Industrial manufacture); PUR
     (Purification or recovery); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)
        (manuf. of D-allo-5-inosose, allo-inositol, D-allo-3-
        inosose, and D-chiro-inositol for pharmaceuticals)
     148218-11-5P, D-Allo-3-Inosose 452335-59-0P,
TT
```

```
RL: BMF (Bioindustrial manufacture); PUR (Purification or recovery); RCT
     (Reactant); BIOL (Biological study); PREP (Preparation); RACT (Reactant or
     reagent)
        (manuf. of D-allo-5-inosose, allo-inositol, D-allo-3-
        inosose, and D-chiro-inositol for pharmaceuticals)
     13762-51-1, Potassium borohydride 16940-17-3, Sodium
     trimethoxyborohydride 16940-66-2, Sodium borohydride
     16949-15-8, Lithium borohydride 25895-60-7, Sodium
     cyanoborohydride
     RL: RCT (Reactant); RACT (Reactant or reagent)
        (reducing agent; manuf. of D-allo-5-inosose, allo-inositol,
        D-allo-3-inosose, and D-chiro-inositol for pharmaceuticals)
L12 ANSWER 3 OF 7 CAPLUS COPYRIGHT 2003 ACS on STN
                         2002:19320 CAPLUS
ACCESSION NUMBER:
DOCUMENT NUMBER:
                         136:68818
                         Microbial manufacture of L-chiro-1-inosose
TITLE:
INVENTOR(S):
                         Takahashi, Atsushi; Kamibe, Kenji;
                         Kita, Yuichi; Mori, Tetsuya;
                         Yamaguchi, Masanori; Tomoda, Akihiro; Tamamura,
                         Takeshi
                         Hokko Chemical Industry Co., Ltd., Japan
PATENT ASSIGNEE(S):
                         Jpn. Kokai Tokkyo Koho, 11 pp.
SOURCE:
                         CODEN: JKXXAF
DOCUMENT TYPE:
                         Patent
                          Japanese
LANGUAGE:
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
     PATENT NO.
                      KIND
                            DATE
                                            APPLICATION NO.
                                                             DATE
     JP 2002000285
                                            JP 2000-186337
                                                             20000621
                       A2
                            20020108
PRIORITY APPLN. INFO.:
                                         JP 2000-186337
                                                             20000621
                         CASREACT 136:68818
OTHER SOURCE(S):
     L-Chiro-1-inosose (I), useful as an enzyme inhibitor or an
     intermediate for pharmaceuticals, is manufd. with microorganisms from
     myo-inositol (II). Xanthomonas sp. AB10198 (FERM P-17893) was cultured in
     a liq. medium contg. II at 27.degree. for 5 days to give I at 141 mg/mL in
     95% conversion.
     87-89-8, myo-Inositol RL: BCP (Biochemical process); RCT (Reactant); BIOL (Biological study);
     PROC (Process); RACT (Reactant or reagent)
        (microbial manuf. of chiro-inosose from myo-inositol)
RN
     87-89-8 CAPLUS
     myo-Inositol (9CI) (CA INDEX NAME)
Relative stereochemistry.
       OH
HO.
             OH
             OH
     56816-02-5P, L-chiro-1-Inosose
     RL: BMF (Bioindustrial manufacture); BPN (Biosynthetic preparation); BIOL
     (Biological study); PREP (Preparation)
        (microbial manuf. of chiro-inosose from myo-inositol)
RN
     56816-02-5 CAPLUS
```

L-chiro-1-Inosose (9CI) (CA INDEX NAME)

Absolute stereochemistry.

D-allo-5-Inosose

```
ОН
             OH
       ÓН
IC
    ICM C12P007-26
         C12N001-20; C12P007-18; C12P007-26; C12R001-64; C12R001-01;
          C12R001-38; C12R001-02; C12R001-18; C12R001-425; C12R001-185;
          C12R001-21
    16-5 (Fermentation and Bioindustrial Chemistry)
    chiroinosose manuf Xanthomonas myoinositol oxidn; microbial
     oxidn inositol inosose manuf
IT
    Oxidation
        (biol.: microbial manuf. of chiro-inosose from myo-inositol)
IT
    Acetobacter
    Acetobacteraceae
    Agrobacterium
    Enterobacter
     Enterobacteriaceae
     Erwinia
     Fermentation
     Gluconobacter
    Haemophilus
     Pasteurella
     Pasteurellaceae
     Pseudomonadaceae
    Pseudomonas
     Rhizobiaceae
     Serratia
     Sphingomonas
     Xanthomonas
     Yersinia
        (microbial manuf. of chiro-inosose from myo-inositol)
    87-89-8, myo-Inositol
     RL: BCP (Biochemical process); RCT (Reactant); BIOL (Biological study);
    PROC (Process); RACT (Reactant or reagent)
        (microbial manuf. of chiro-inosose from myo-inositol)
     56816-02-5P, L-chiro-1-Inosose
     RL: BMF (Bioindustrial manufacture); BPN (Biosynthetic preparation); BIOL
     (Biological study); PREP (Preparation)
(microbial manuf. of chiro-inosose from myo-inositol)
L12 ANSWER 4 OF 7 CAPLUS COPYRIGHT 2003 ACS on STN
ACCESSION NUMBER:
                         2001:874389 CAPLUS
DOCUMENT NUMBER:
                          136:20217
                         Preparation of L-epi-inositol
TITLE:
INVENTOR(S):
                         Ogawa, Seiichiro; Takahashi, Atsushi
                         Hokko Chemical Industry Co., Ltd., Japan
PATENT ASSIGNEE(S):
SOURCE:
                          Jpn. Kokai Tokkyo Koho, 22 pp.
                          CODEN: JKXXAF
DOCUMENT TYPE:
                         Patent
LANGUAGE:
                          Japanese
FAMILY ACC. NUM. COUNT:
                         1
PATENT INFORMATION:
     PATENT NO.
                      KIND
                             DATE
                                            APPLICATION NO.
     JP 2001335544
                             20011204
                                            JP 2000-158238
                                                              20000529
                       A2
PRIORITY APPLN. INFO.:
                                         JP 2000-158238
                                                              20000529
OTHER SOURCE(S):
                         CASREACT 136:20217; MARPAT 136:20217
```

AB Title compds. I (R1, R4-R7 = H, acyl, alkyl; if R2 = amino, acylamido, alkylamino, N-acyl-N-alkylamino, then R3 = H; if R2 = OH, acyloxy, then R3 = HOCH2, acyloxymethyl, azidomethyl, aminomethyl, acylamidomethyl; if R2 = O, then R3 = CH2 forming ring with R2) or their salts are prepd. L-Epi-2-inosose was reacted with PhNHNH2 in the presence of AcOH in H2O at 5.degree. for 2 h to give 79.5% L-epi-2-inosose phenylhydrazone, which was hydrogenated with H in the presence of platinum oxide in AcOH, treated with HCl at 100.degree. for 3.5 h, and treated with strongly acidic ion exchanger to give 2-amino-2-deoxy-L-epi-inositol.

RN 377777-76-9 CAPLUS

CN D-epi-Inositol, 4-C-(azidomethyl)-, 1,2,3,5,6-pentaacetate (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

IT 22059-57-0P 38876-94-7P 377777-72-5P 377777-74-7P 377777-77-0P 379224-06-3P 379224-11-0P

RL: IMF (Industrial manufacture); SPN (Synthetic preparation); PREP (Preparation)

(prepn. of epi-inositol)

RN 22059-57-0 CAPLUS

CN D-epi-Inositol, 2,21-anhydro-2-C-(hydroxymethyl)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RN 38876-94-7 CAPLUS
CN D-epi-Inositol, 4-C-[(acetyloxy)methyl]-, 1,2,3,5,6-pentaacetate (9CI)
(CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RN 377777-72-5 CAPLUS

CN D-epi-Inositol, 4-deoxy-4-(ethylamino)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RN 377777-74-7 CAPLUS

CN D-epi-Inositol, 4-deoxy-4-(propylamino)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RN 377777-77-0 CAPLUS

CN D-epi-Inositol, 4-C-(azidomethyl)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 379224-06-3 CAPLUS

CN D-epi-Inositol, 4-amino-4-deoxy- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

RN 379224-11-0 CAPLUS CN 1-0xaspiro[2.5]octane-4,5,6,7,8-pentol, pentaacetate, (4R,55,75,8S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

T79-03-8, Propionyl chloride 100-63-0, Phenylhydrazine
334-88-3, Diazomethane 33471-33-9, D-epi-4Inosose
RI: RCT (Reactant): RACT (Reactant or reagent)

RL: RCT (Reactant); RACT (Reactant or reagent)
 (prepn. of epi-inositol)

RN 79-03-8 CAPLUS

CN Propanoyl chloride (9CI) (CA INDEX NAME)

RN 100-63-0 CAPLUS CN Hydrazine, phenyl- (8CI, 9CI) (CA INDEX NAME)

H₂N--NH--Ph

RN 334-88-3 CAPLUS CN Methane, diazo- (8CI, 9CI) (CA INDEX NAME)

 $H_2C = N \stackrel{+}{=} N^-$

RN 33471-33-9 CAPLUS CN D-epi-4-Inosose (9CI) (CA INDEX NAME)

Absolute stereochemistry.

IT 7045-49-0P 377777-73-6P 377777-75-8P

379224-07-4P 379224-09-6P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(prepn. of epi-inositol)

RN 7045-49-0 CAPLUS

CN D-epi-4-Inosose, phenylhydrazone (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 377777-73-6 CAPLUS

CN D-epi-Inositol, 4-(acetylethylamino)-4-deoxy-, pentaacetate (ester) (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

RN 377777-75-8 CAPLUS

CN D-epi-Inositol, 4-(acetylpropylamino)-4-deoxy-, pentaacetate (ester) (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RN 379224-07-4 CAPLUS

CN D-epi-Inositol, 4-(acetylamino)-4-deoxy-, 1,2,3,5,6-pentaacetate (9CI)
 (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

379224-09-6 CAPLUS

D-epi-Inositol, 4-amino-4-deoxy-, hydrochloride (9CI) (CA INDEX NAME) CN

Absolute stereochemistry.

HC1

```
ICM C07C213-02
IC
        C07C067-26; C07C069-21; C07C215-44; C07C233-23; C07C247-06;
         CO7D301-02; CO7D303-14; CO7B061-00
```

- CC 33-6 (Carbohydrates)
- ST inositol prepn
- 377777-76-9P

RL: IMF (Industria) manufacture); RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent) (prepn. of epi-inositol)

22059-57-0P 38876-94-7P 377777-72-5P 377777-74-7P 377777-77-0P 379224-06-3P

379224-11-0P

RL: IMF (Industrial manufacture); SPN (Synthetic preparation); PREP (Preparation)

(prepn. of epi-inositol)

79-03-8, Propionyl chloride 100-63-0, Phenylhydrazine 334-88-3, Diazomethane 33471-33-9, D-epi-4-

Triosose

RL: RCT (Reactant); RACT (Reactant or reagent)

(prepn. of epi-inositol)

7045-49-0P 377777-73-6P 377777-75-8P

379224-07-4P 379224-09-6P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent) (prepn. of epi-inositol)

L12 ANSWER 5 OF 7 CAPLUS COPYRIGHT 2003 ACS on STN

ACCESSION NUMBER:

2001:798044 CAPLUS

DOCUMENT NUMBER: TITLE:

135:339209

Compositions for inhibiting the proliferation of human

immunodeficiency virus and method of inhibiting the proliferation of this virus

Takeuchi, Tomio; Ohno, Tuneya; Nakamura, INVENTOR(S):

Mariko: Tamamura, Tsuyoshi; Takahashi,

Atsushi

PATENT ASSIGNEE(S): Hokko Chemical Industry Co., Ltd., Japan; Zaidan Hojin Biseibutsu Kagaku Kenkyu Kai

SOURCE: PCT Int. Appl., 27 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

Japanese

FAMILY ACC. NUM. COUNT:

PATENT INFORMATION:

PATENT NO. KIND DATE APPLICATION NO. DATE
WO 2001080848 A1 20011101 WO 2001-JP3587 20010425

W: CA, JP, US

RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,

PT, SE, TR

PRIORITY APPLN. INFO.: JP 2000-123407 A 20000425

AB (+)-Protoquercitol, (-)-protoquercitol, (.+-.)-protoquercitol, L-epi-2-inosose, D-epi-2-inosose, and DL-epi-2-inosose

have an activity of inhibiting the proliferation of HIV infecting human T cells and/or human monocytes/macrophages and/or other human hemocytes and, therefore, are useful as HIV proliferation inhibitors. Also, a method of inhibiting the proliferation of HIV by treating HIV with the above compds. or enantiomers or racemates thereof, is provided.

IT 488-68-6, D-epi-2-Inosose 488-73-3,

(+)-Proto-quercitol 6623-68-3, DL-epi-2-Inosose

17278-12-5 33471-33-9, 2-Inosose, L-epi-

90899-07-3

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(quercitol and inosose analogs for inhibition of HIV proliferation)

RN 488-68-6 CAPLUS

CN D-epi-2-Inosose (7CI, 9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 488-73-3 CAPLUS

N D-chiro-Inositol, 2-deoxy- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (+).

RN 6623-68-3 CAPLUS

CN epi-2-Inosose (9CI) (CA INDEX NAME)

Relative stereochemistry.

RN 17278-12-5 CAPLUS

CN L-chiro-Inositol, 2-deoxy- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

RN 33471-33-9 CAPLUS

CN D-epi-4-Inosose (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 90899-07-3 CAPLUS

CN chiro-Inositol, 2-deoxy- (9CI) (CA INDEX NAME)

Relative stereochemistry.

- IC ICM A61K031-047
 - ICS A61K031-122; A61P031-18
- CC 1-5 (Pharmacology)
 - Section cross-reference(s): 63
- ST quercitol inosose HIV proliferation inhibitor
- IT Hemocyte
 - Macrophage
 - Monocyte
 - T cell (lymphocyte)

(infection; quercitol and inosose analogs for inhibition of

HIV proliferation)

IT Drug delivery systems

(injections; quercitol and inosose analogs for inhibition of HIV proliferation)

IT Anti-AIDS agents

Human immunodeficiency virus 1 (quercitol and inosose analogs for inhibition of HIV proliferation) IT 488-68-6, D-epi-2-Inosose 488-73-3, (+)-Proto-quercitol 6623-68-3, DL-epi-2-Inosose 17278-12-5 33471-33-9, 2-Inosose, L-epi-90899-07-3 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (quercitol and inosose analogs for inhibition of HIV proliferation) REFERENCE COUNT: q THERE ARE 9 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT L12 ANSWER 6 OF 7 CAPLUS COPYRIGHT 2003 ACS on STN ACCESSION NUMBER: 2001:24366 CAPLUS DOCUMENT NUMBER: 134:171044 TITLE: (-)-epi-Inosose-2 AUTHOR(S): Hosomi, Hiroyuki; Ohba, Shigeru; Ogawa, Seiichiro; Takahashi, Atsushi Faculty of Science and Technology, Department of CORPORATE SOURCE: Chemistry, Keio University, Kohoku-ku, Yokohama, 223-8522, Japan Acta Crystallographica, Section C: Crystal Structure Communications (2000), C56(12), e584-e585 CODEN: ACSCEE; ISSN: 0108-2701 SOURCE: Munksgaard International Publishers Ltd. PUBLISHER: DOCUMENT TYPE: Journal LANGUAGE: English The structure of the title compd., C6H1006, was detd. to confirm the position of the keto group in the mol. prepd. enantioselectively by a bioconversion from myo-inositol. There are two independent mols. showing similar geometry. Crystallog, data are given. 33471-33-9, (-)-Epi-Inosose-2 TT RL: PRP (Properties) (crystal structure of) RN 33471-33-9 CAPLUS D-epi-4-Inosose (9CI) (CA INDEX NAME) Absolute stereochemistry. HO OH ОН ОН 75-8 (Crystallography and Liquid Crystals) Section cross-reference(s): 33 mol structure epi inosose Crystal structure TT Molecular structure (of epi-inosose-2) 33471-33-9, (-)-Epi-Inosose-2 RL: PRP (Properties) (crystal structure of)

L12 ANSWER 7 OF 7 CAPLUS COPYRIGHT 2003 ACS on STN

11

ACCESSION NUMBER: 2000:881342 CAPLUS

DOCUMENT NUMBER: 134:42384

REFERENCE COUNT:

THERE ARE 11 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

```
Novel process for producing L-epi-2-inosose
TITLE:
                           by microbial oxidation of myo-inositol and novel
                           process for producing epi-inositol
INVENTOR(S):
                           Takahashi, Atsushi; Kanbe, Kenji;
                           Mori, Tetsuya; Kita, Yuichi;
                           Tamamura, Tsuyoshi; Takeuchi, Tomio
                           Hokko Chemical Industry Co., Ltd., Japan; Zaidan Hojin
PATENT ASSIGNEE(S):
                           Biseibutsu Kagaku Kenkyu Kai
                           PCT Int. Appl., 65 pp.
SOURCE:
                           CODEN: PIXXD2
DOCUMENT TYPE:
                           Patent
LANGUAGE:
                           Japanese
FAMILY ACC. NUM. COUNT:
PATENT INFORMATION:
                                               APPLICATION NO.
                                                                 DATE
     PATENT NO.
                       KIND DATE
                                                                 20000607
                                               WO 2000-JP3687
     WO 2000075355
                        A1
                              20001214
         W: CA, CN, IL, IN, JP, KR, US
         RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
              PT, SE
                                               EP 2000-937174
                                                                 20000607
     EP 1197562
                              20020417
                         A1
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
              IE, SI, LT, LV, FI, RO
                                                                19990607
PRIORITY APPLN. INFO.:
                                           JP 1999-159861
                                                             Α
                                           JP 1999-340523
                                                             Α
                                                                 19991130
                                           JP 2000-151709
                                                                 20000523
                                           WO 2000-JP3687
                                                                 20000607
OTHER SOURCE(S):
                           CASREACT 134:42384
     L-Epi-2-inosose and epi-inositol, which are
     useful as various drugs or synthesis intermediates, can be efficiently
     produced from less expensive myo-inositol. Myo-inositol is treated with a
     gram-neg. bacterium. e.g. Xanthomonas sp., capable of converting
     myo-inositol into L-epi-2-inosose to thereby convert the
     \label{eq:myo-inositol} \mbox{ into $L$-epi-$2$-inosose.} \quad \mbox{The $L$-epi-$2$-inosose}
     thus obtained is further reacted in an aq. reaction medium with a reducing agent comprising an alkali metal boron hydride or another alkali metal
     hydride to form epi-inositol and myo-inositol. Next,
     the epi-inositol is sepd. and isolated from the redn.
     reaction mixt. comprising epi-inositol and myo-inositol to give epi-inositol.
     6623-68-3P, epi-2-Inosose
     RL: BPN (Biosynthetic preparation); RCT (Reactant); BIOL (Biological
     study); PREP (Preparation); RACT (Reactant or reagent)
        (novel process for producing L-epiinosose by microbial oxidn.
        of myo-inositol and boron hydride-redn. to epi-
        inositol)
     6623-68-3 CAPLUS
RN
     epi-2-Inosose (9CI) (CA INDEX NAME)
CN
Relative stereochemistry.
       OH
              OH
HO.
       R
     R
       OH
```

```
RN 87-89-8 CAPLUS
CN myo-Inositol (9CI) (CA INDEX NAME)
```

Relative stereochemistry.

Relative stereochemistry.

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C12P019-02; C12N001-20; C12P019-02; C12R001-64; C12P019-02; C12R001-38; C12P019-02; C12R001-02; C12P019-02; C12R001-18; C12P019-02; C12R001-425; C12P019-02; C12R001-21; C12P019-02; C12R001-01; C12N001-20; C12R001-64;
     C12N001-20; C12R001-38
     33-6 (Carbohydrates)
CC
      Section cross-reference(s): 16
     gram neg bacterium Xanthomonas microbial oxidn myoinositol;
     epiinosose prepn redn; epiinositol prepn
TT
     Oxidation
         (biol.; novel process for producing L-epiinosose by microbial
         oxidn. of myo-inositol and boron hydride-redn. to epi-
         inositol)
     Acetobacter
     Acetobacteraceae
     Agrobacterium
     Enterobacter
      Enterobacteriaceae
      Erwinia
     Gluconobacter
     Gram-negative bacteria
     Haemophilus
     Pasteurella
      Pasteurellaceae
      Pseudomonadaceae
      Pseudomonas
     Reduction
     Rhizobiaceae
      Serratia
      Xanthomonas
      Yersinia
         (novel process for producing L-epiinosose by microbial oxidn.
         of myo-inositol and boron hydride-redn. to epi-
```

inositol)

6623-68-3P, epi-2-Inosose

RL: BPN (Biosynthetic preparation); RCT (Reactant); BIOL (Biological

study); PREP (Preparation); RACT (Reactant or reagent) (novel process for producing L-epiinosose by microbial oxidn.

of myo-inositol and boron hydride-redn. to epiinositol)

IT

87-89-8, myo-Inositol RL: RCT (Reactant); RACT (Reactant or reagent)

(novel process for producing L-epiinosose by microbial oxidn.

of myo-inositol and boron hydride-redn. to epiinositol)

IT

488-58-4P, epi-Inositol RL: SPN (Synthetic preparation); PREP (Preparation)

(novel process for producing L-epiinosose by microbial oxidn.

of myo-inositol and boron hydride-redn. to epi-

inositol)

REFERENCE COUNT:

THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT